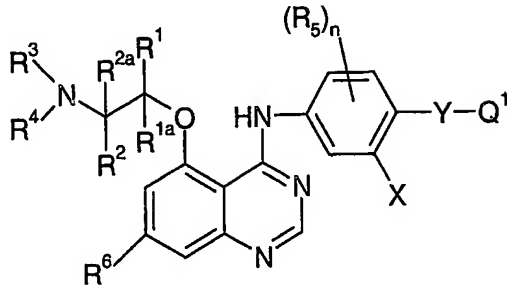
CLAIMS

1. A quinazoline derivative of the formula I:



I

wherein:

each of R^1 and R^2, which may be the same or different, is selected from hydrogen, carboxy, cyano, formyl, (1-3C)alkyl, (2-3C)alkanoyl, (1-3C)alkoxycarbonyl, carbamoyl, N-(1-3C)alkylcarbamoyl and N, N-di-[(1-3C)alkyl]carbamoyl;

each of R^{1a} and R^{2a}, which may be the same or different, is selected from hydrogen and (1-3C)alkyl;

each of R^3 and R^4, which may be the same or different, is selected from hydrogen, (1-3C)alkyl and (2-4C) alkenyl;

and wherein any CH or CH_2 or CH_3 within any of R^1, R^{1a}, R^2, R^{2a}, R^3 and R^4 optionally bears on each said CH or CH_2 or CH_3 one or more (for example 1, 2 or 3) halogeno substituents or a substituent selected from hydroxy, cyano, (1-3C)alkoxy, amino, (2-3C)alkanoyl, (1-3C)alkylamino and di-[(1-3C)alkyl]amino;

X is selected from hydrogen, halogeno, (1-4C)alkyl, (1-4C)alkoxy, (2-4C)alkenyl and (2-4C)alkynyl;

each R^5, which may be the same or different, is selected from halogeno, hydroxy, (1-4C)alkyl, (1-4C)alkoxy, (2-4C)alkenyl and (2-4C)alkynyl;

Y is selected from a direct bond, O, S, $OC(R^7)_2$, $SC(R^7)_2$, SO, SO_2, $N(R^7)$, CO and $N(R^7)C(R^7)_2$ wherein each R^7 is, independently, hydrogen or (1-6C)alkyl;

Q^1 is selected from phenyl, pyridyl, pyrazinyl, 1,3-thiazolyl, 1H-imidazolyl, 1H-pyrazolyl, 1,3-oxazolyl and isoxazolyl,

and wherein Q^1 optionally bears one or more substituents (for example 1, 2 or 3), which may be the same or different, selected from halogeno, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, sulfamoyl, formyl, mercapto, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl,

(1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, (3-6C)alkenoylamino, N-(1-6C)alkyl-(3-6C)alkenoylamino, (3-6C)alkynoylamino, N-(1-6C)alkyl-(3-6C)alkynoylamino, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino, and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, or from a group of the formula:

$$-X^1-R^8$$

wherein X^1 is a direct bond or is selected from O, CO and $N(R^9)$, wherein R^9 is hydrogen or (1-6C)alkyl, and R^8 is halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, carboxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, N-(1-6C)alkylamino-(1-6C)alkyl, N,N-di-[(1-6C)alkyl]amino-(1-6C)alkyl, (2-6C)alkanoylamino-(1-6C)alkyl, (1-6C)alkoxycarbonylamino-(1-6C)alkyl, carbamoyl-(1-6C)alkyl, N-(1-6C)alkylcarbamoyl-(1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkyl, (2-6C)alkanoyl-(1-6C)alkyl or (1-6C)alkoxycarbonyl-(1-6C)alkyl,

and wherein any CH_2 or CH_3 within a substituent on Q^1 optionally bears on each said CH_2 or CH_3 one oR more (for example 1, 2, or 3) halogeno or (1-6C)alkyl substituents or a substituent selected from hydroxy, cyano, amino, (1-4C)alkoxy, (1-4C)alkylamino and di-[(1-4C)alkyl]amino;

$\mathbf{R^6}$ is selected from hydrogen, (1-6C)alkoxy, (2-6C)alkenyloxy and (2-6C)alkynyloxy,

and wherein any CH_2 or CH_3 group within a R^6 substituent optionally bears on each said CH_2 or CH_3 group one or more halogeno or (1-6C)alkyl substituents, or a substituent selected from hydroxy and (1-6C)alkoxy;

n is 0, 1, 2 or 3;

or a pharmaceutically acceptable salt thereof.

2. A quinazoline derivative of the formula I as defined in claim 1, wherein R^1 is selected from hydrogen, methyl and ethyl, R^2 is selected from hydrogen, carboxy, cyano, methyl, ethyl, acetyl, methoxycarbonyl, carbamoyl, N-methylcarbamoyl and N, N-di-methylcarbamoyl, and R^{1a} and R^{2a} are each hydrogen.

3. A quinazoline derivative of the formula I as defined in claim 1, wherein R^2 is selected from hydrogen, methyl and ethyl, R^1 is selected from hydrogen, carboxy, cyano, methyl, ethyl, acetyl, methoxycarbonyl, carbamoyl, N-methylcarbamoyl and N, N-di-methylcarbamoyl, and R^{1a} and R^{2a} are each hydrogen.

4. A quinazoline derivative of the formula I as defined in claim 1, wherein R^1 and R^{1a} are each hydrogen, R^2 is selected from hydrogen, carboxy, cyano, methyl, ethyl, acetyl, methoxycarbonyl, carbamoyl, N-methylcarbamoyl and N, N-di-methylcarbamoyl, and R^{2a} is selected from hydrogen and (1-3C)alkyl.

5. A quinazoline derivative of the formula I as defined in claim 1, wherein R^2 and R^{2a} are each hydrogen, R^1 is selected from hydrogen, carboxy, cyano, methyl, ethyl, acetyl, methoxycarbonyl, carbamoyl, N-methylcarbamoyl and N, N-di-methylcarbamoyl, and R^{1a} is selected from hydrogen and (1-3C)alkyl.

6. A quinazoline derivative of the formula I as defined in any one of claims 1, 2, 3 and 5, wherein R^1 is methyl, and R^2, R^{1a} and R^{2a} are each hydrogen.

7. A quinazoline derivative of the formula I as defined in any one of claims 1 to 4, wherein R^2 is methyl and R^1, R^{1a} and R^{2a} are each hydrogen.

8. A quinazoline derivative of the formula I as defined in claim 1 or claim 5, wherein R^1 and R^{1a} are each methyl and R^2 and R^{2a} are each hydrogen.

9. A quinazoline derivative of the formula I as defined in claim 1 or claim 4, wherein R^2 and R^{2a} are each methyl and R^1 and R^{1a} are each hydrogen.

10. A quinazoline derivative of the formula I as defined in any one of the preceding claims, wherein each of R^3 and R^4, which may be the same or different, is selected from (1-3C)alkyl, wherein any CH or CH_2 or CH_3 within any of R^3 and R^4 optionally bears on each said CH or CH_2 or CH_3 one or more substituents selected from hydroxy and (1-3C)alkoxy.

11. A quinazoline derivative of the formula I as defined in any one of claims 1 to 9, wherein each of R^3 and R^4, which may be the same or different, is selected from hydrogen, methyl, ethyl, propenyl, 2-methoxyethyl and 2-hydroxyethyl.

12. A quinazoline derivative of the formula I as defined in claim 11, wherein each of R^3 and R^4, which may be the same or different, is selected from methyl, ethyl, propenyl, 2-methoxyethyl and 2-hydroxyethyl.

13. A quinazoline derivative of the formula I as defined in claim 11 or claim 12, wherein R^3 is methyl and R^4 is selected from methyl, ethyl, 2-hydroxyethyl, 2-methoxyethyl and propenyl.

14. A quinazoline derivative of the formula I as defined in any one of claims 10 to 13, wherein R^3 and R^4 are each methyl.

15. A quinazoline derivative of the formula I as defined in any one of claims 10 to 12, wherein R^3 is ethyl and R^4 is 2-hydroxyethyl.

16. A quinazoline derivative of the formula I as defined in any one of the preceding claims, wherein X is selected from hydrogen, halogeno, (1-4C)alkyl and (1-4C)alkoxy.

17. A quinazoline derivative of the formula I as defined in claim 16, wherein X is selected from hydrogen, fluoro, chloro, methyl and methoxy.

18. A quinazoline derivative of the formula I as defined in claim 16 or claim 17, wherein X is selected from methyl and chloro.

19. A quinazoline derivative of the formula I as defined in claim 18, wherein X is chloro.

20. A quinazoline derivative of the formula I as defined in claim 18, wherein X is methyl.

21. A quinazoline derivative of the formula I as defined in any one of the preceding claims, wherein Y is selected from O, S and $OC(R^7)_2$ wherein each R^7 is, independently, hydrogen or (1-4C)alkyl.

22. A quinazoline derivative of the formula I as defined in claim 21, wherein Y is selected from O, S and OCH_2.

23. A quinazoline derivative of the formula I as defined in claim 21 or claim 22, wherein Y is O.

24. A quinazoline derivative of the formula I as defined in claim 21 or claim 22, wherein Y is S.

25. A quinazoline derivative of the formula I as defined in claim 21 or claim 22, wherein Y is OCH_2.

26. A quinazoline derivative of the formula I as defined in any one of the preceding claims, wherein n is 0.

27. A quinazoline derivative of the formula I as defined in any one of the preceding claims, wherein Q^1 is selected from phenyl, 2-pyridyl, 2-pyrazinyl, 1,3-thiazol-4-yl, 1,3-thiazol-5-yl, 1H-imidazol-2-yl and isoxazol-3-yl, and wherein Q^1 optionally bears one or more substituents, which may be the same or different, as defined in claim 1.

28. A quinazoline derivative of the formula I as defined in claim 27, wherein Q^1 is selected from phenyl, 2-pyridyl, 2-pyrazinyl, 1,3-thiazol-4-yl, 1,3-thiazol-5-yl, 1H-imidazol-2-yl and 3-isoxazolyl, and wherein Q^1 optionally bears one or more substituents, which may be the same or different, selected from fluoro and (1-4C)alkyl.

29. A quinazoline derivative of the formula I as defined in claim 27 or claim 28, wherein Q^1 is selected from 3-fluorophenyl, 2-pyridyl, 2-pyrazinyl, 1-methyl-1H-imidazol-2-yl, 1,3-thiazol-4-yl, 1,3-thiazol-5-yl and 5-methyl-3-isoxazolyl.

30. A quinazoline derivative of the formula I as defined in any one of the preceding claims, wherein R^6 is hydrogen.

31. A quinazoline derivative selected from one or more of the following:

4-(3-Chloro-4-(2-pyridylmethoxy)anilino)-5-(2dimethylaminoethoxy)quinazoline;

4-(3-Chloro-4-(2-pyridylmethoxy)anilino)-5-(2-dimethylamino-1-methylethoxy)quinazoline;

4-(3-Chloro-4-(1-methyl-1*H*-imidazol-2-ylthio)anilino)-5-(2-dimethylaminoethoxy)quinazoline;

4-(3-Chloro-4-(1-methyl-1*H*-imidazol-2-ylthio)anilino)-5-(2-dimethylamino-2-methylethoxy)quinazoline;

4-(4-(3-Fluorobenzyloxy)anilino)-5-(2-dimethylaminoethoxy)quinazoline;

4-(4-(3-Fluorobenzyloxy)anilino)-5-(2-dimethylamino-1-methylethoxy)quinazoline;

4-(3-Chloro-4-(2-pyrazinylmethoxy)anilino)-5-(2-dimethylaminoethoxy)quinazoline;

4-(3-Chloro-4-(2-pyrazinylmethoxy)anilino)-5-(2-dimethylamino-1-methylethoxy)quinazoline;

4-(3-Chloro-4-(5-methylisoxazol-3-ylmethoxy)anilino)-5-(2-dimethylaminoethoxy)quinazoline;

4-(3-Chloro-4-(5-methylisoxazol-3-ylmethoxy)anilino)-5-(2-dimethylamino-1-methylethoxy)quinazoline;

4-(3-Chloro-4-(3-fluorobenzyloxy)anilino)-5-(2-(N-ethyl-N-methylamino)ethoxy)quinazoline;

4-(3-Chloro-4-(3-fluorobenzyloxy)anilino)-5-(2-dimethylaminoethoxy)quinazoline;

4-(3-Chloro-4-(3-fluorobenzyloxy)anilino)-5-[2-(N-(2-hydroxyethyl)-N-methylamino)ethoxy]quinazoline;

4-(3-Chloro-4-(2-pyridylmethoxy)anilino)- 5-(2-(N-ethyl-N-methylamino)ethoxy)quinazoline;

4-(3-Chloro-4-(2-pyridylmethoxy)anilino)- 5-(2-(N-(2-hydroxyethyl)-N-methylamino)ethoxy)quinazoline;

4-(3-Chloro-4-(3-fluorobenzyloxy)anilino)-5-(2-dimethylamino-2-methylethoxy)quinazoline;

4-(3-Chloro-4-(2-pyridylmethoxy)anilino)-5-(2-dimethylamino-2-methylethoxy)quinazoline;

N-[3-Chloro-4-(1,3-thiazol-4-ylmethoxy)phenyl]-5-[2-(dimethylamino)ethoxy]quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-yloxy)phenyl]-5-[2-(dimethylamino)ethoxy]quinazolin-4-amine;

N-[3-Chloro-4-(1,3-thiazol-4-ylmethoxy)phenyl]-5-[(1*R*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-[3-Chloro-4-(pyrazin-2-ylmethoxy)phenyl]-5-[(1*R*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-{3-Chloro-4-[(3-fluorobenzyl)oxy]phenyl}-5-[(1*R*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-[2-(dimethylamino)-2-methylpropoxy]quinazolin-4-amine;

N-[3-Chloro-4-(1,3-thiazol-4-ylmethoxy)phenyl]-5-[2-(dimethylamino)-2-methylpropoxy]quinazolin-4-amine;

N-{3-Chloro-4-[(5-methylisoxazol-3-yl)methoxy]phenyl}-5-[2-(dimethylamino)-2-methylpropoxy]quinazolin-4-amine;

5-[2-(Dimethylamino)ethoxy]-*N*-[3-methyl-4-(pyridin-2-ylmethoxy)phenyl]quinazolin-4-amine;

5-[2-(Dimethylamino)ethoxy]-*N*-[3-methyl-4-(1,3-thiazol-4-ylmethoxy)phenyl]quinazolin-4-amine;

5-[2-(Dimethylamino)ethoxy]-*N*-{3-methyl-4-[(5-methylisoxazol-3-yl)methoxy]phenyl}quinazolin-4-amine;

5-[(1*R*)-2-(Dimethylamino)-1-methylethoxy]-*N*-[3-methyl-4-(pyridin-2-ylmethoxy)phenyl]quinazolin-4-amine;

5-[(1*R*)-2-(Dimethylamino)-1-methylethoxy]-*N*-[3-methyl-4-(pyrazin-2-ylmethoxy)phenyl]quinazolin-4-amine;

5-[(1*R*)-2-(dimethylamino)-1-methylethoxy]-*N*-[3-methyl-4-(1,3-thiazol-4-ylmethoxy)phenyl]quinazolin-4-amine;

5-[(1*R*)-2-(Dimethylamino)-1-methylethoxy]-*N*-{3-methyl-4-[(5-methylisoxazol-3-yl)methoxy]phenyl}quinazolin-4-amine;

N-[3-Chloro-4-(pyrazin-2-ylmethoxy)phenyl]-5-[(1*S*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-{3-Chloro-4-[(3-fluorobenzyl)oxy]phenyl}-5-[(1*S*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-[(1*R*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-(3-Chloro-4-{[3-(trifluoromethyl)benzyl]oxy}phenyl)-5-[(1*R*)-2-(dimethylamino)-1-

5-[2-(dimethylamino)-2-methylpropoxy]-*N*-[3-methyl-4-(1,3-thiazol-4-ylmethoxy)phenyl]quinazolin-4-amine;

5-[2-(Dimethylamino)ethoxy]-*N*-{3-methoxy-4-[(5-methylisoxazol-3-yl)methoxy]phenyl}quinazolin-4-amine;

5-[2-(Dimethylamino)ethoxy]-*N*-[3-methoxy-4-(pyrazin-2-ylmethoxy)phenyl]quinazolin-4-amine;

5-[2-(Dimethylamino)ethoxy]-*N*-[3-fluoro-4-(1,3-thiazol-5-ylmethoxy)phenyl]quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-[(1*S*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-{[(2*S*)-2-(dimethylamino)propyl]oxy}quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-{[(2*R*)-2-(dimethylamino)propyl]oxy}quinazolin-4-amine;

5-{2-[Allyl(methyl)amino]ethoxy}-*N*-[3-chloro-4-(pyridin-2-ylmethoxy)phenyl]quinazolin-4-amine;

2-[{2-[(4-{[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]amino}quinazolin-5-yl)oxy]ethyl}(ethyl)amino]ethanol;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-{(1*S*)-2-[(2-methoxyethyl)(methyl)amino]-1-methylethoxy}quinazolin-4-amine;

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-{(1*R*)-2-[ethyl(methyl)amino]-1-methylethoxy}quinazolin-4-amine;

5-{(1*R*)-2-[Allyl(methyl)amino]-1-methylethoxy}-*N*-[3-chloro-4-(pyridin-2-ylmethoxy)phenyl]quinazolin-4-amine;

5-{(1*S*)-2-[Allyl(methyl)amino]-1-methylethoxy}-*N*-[3-chloro-4-(pyridin-2-ylmethoxy)phenyl]quinazolin-4-amine;

N-{3-Chloro-4-[(3-fluorobenzyl)oxy]phenyl}-5-{[(2*S*)-2-(dimethylamino)propyl]oxy}quinazolin-4-amine;

N-{3-Chloro-4-[(3-fluorobenzyl)oxy]phenyl}-5-{[(2*R*)-2-(dimethylamino)propyl]oxy}quinazolin-4-amine;

N-{3-Chloro-4-[(1-methyl-1*H*-imidazol-2-yl)thio]phenyl}-5-{[(2*S*)-2-(dimethylamino)propyl]oxy}quinazolin-4-amine;

N-{3-Chloro-4-[(1-methyl-1*H*-imidazol-2-yl)thio]phenyl}-5-{[(2*R*)-2-(dimethylamino)propyl]oxy}quinazolin-4-amine;

N-{3-Chloro-4-[(1-methyl-1*H*-imidazol-2-yl)thio]phenyl}-5-[(1*R*)-2-(dimethylamino)-1-methylethoxy]quinazolin-4-amine;

5-[2-(Dimethylamino)-1-methylethoxy]-*N*-(3-methoxy-4-phenoxyphenyl)quinazolin-4-amine;

5-[2-(Dimethylamino)-1-methylethoxy]-N-(3-methoxy-4-phenoxyphenyl)quinazolin-4-amine; and

N-[3-Chloro-4-(pyridin-2-ylmethoxy)phenyl]-5-[2-(dimethylamino)-1,1-dimethylethoxy]quinazolin-4-amine;

or a pharmaceutically acceptable salt thereof.

32. A pharmaceutical composition which comprises a quinazoline derivative of the formula I, or a pharmaceutically acceptable salt thereof, as defined in any one of claims 1 to 31 in association with a pharmaceutically-acceptable diluent or carrier.

33. A quinazoline derivative of the formula I, or a pharmaceutically-acceptable salt thereof, as defined in any one of claims 1 to 31 for use as a medicament.

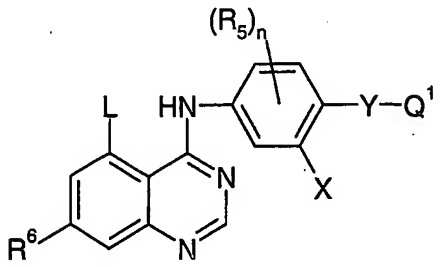
34. A quinazoline derivative of the formula I, or a pharmaceutically acceptable salt thereof, as defined in any one of claims 1 to 31 for use in the production of an anti-proliferative effect which effect is produced alone or in part by inhibiting erbB2 receptor tyrosine kinase in a warm-blooded animal such as man.

35. A quinazoline derivative of the formula I, or a pharmaceutically acceptable salt thereof, as defined in any one of claims 1 to 31 for use in the production of an erbB2 receptor tyrosine kinase inhibitory effect in a warm-blooded animal such as man.

36. A quinazoline derivative of the formula I, or a pharmaceutically acceptable salt thereof, as defined in any one of claims 1 to 31 for use in the production of a selective erbB2 receptor tyrosine kinase inhibitory effect in a warm-blooded animal such as man.

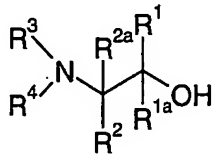
37. A process for the preparation of a quinazoline derivative of the formula I, or a pharmaceutically acceptable salt thereof, as defined in claim 1 which comprises:

(a) the reaction, conveniently in the presence of a suitable base, of a quinazoline of the formula **II**:



II

wherein R^5, R^6, Q^1, X, Y and n are as defined in claim 1 except that any functional group is protected if necessary, and L is a displaceable group, with an alcohol of the formula **III**

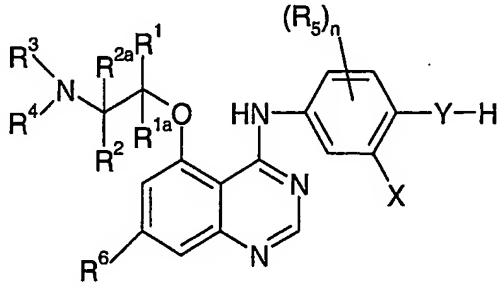


III

wherein R^1, R^{1a}, R^2, R^{2a}, R^3 and R^4 are as defined in claim 1 except that any functional group is protected if necessary;

or

(b) for the preparation of those compounds of the formula **I** wherein Y is $OC(R^7)_2$, $SC(R^7)_2$ or $N(R^7)C(R^7)_2$, the reaction, conveniently in the presence of a suitable base, of a quinazoline of the formula **IV**:



IV

wherein Y is O, S or $N(R^7)$, and X, R^1, R^{1a}, R^2, R^{2a}, R^3, R^4, R^5, R^6, R^7 and n are as defined in claim 1 except that any functional group is protected if necessary, with a compound of the formula **V**:

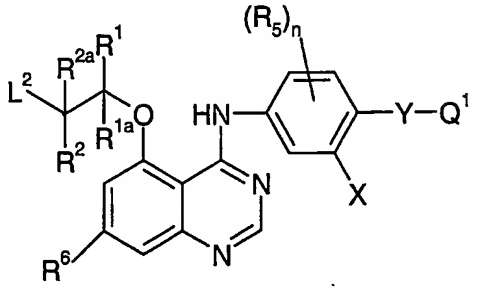
$$Q^1\text{-}C(R^7)_2\text{-}L^1$$

V

wherein L^1 is a suitable displaceable group and Q^1 and R^7 are as defined in claim 1 except that any functional group is protected if necessary;

or

(c) the reaction of a quinazoline of the formula VI:



VI

wherein L^2 is a suitable displaceable group and Q^1, X, Y, R^1, R^{1a}, R^2, R^{2a}, R^5, R^6 and n are as defined in claim 1 except that any functional group is protected if necessary, with an amine of the formula VII:

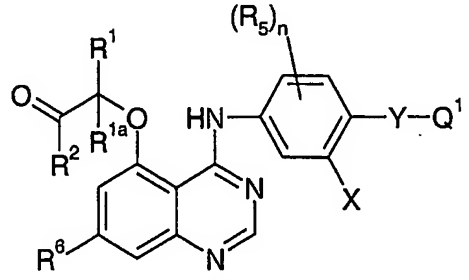
$$NHR^3R^4$$

VII

wherein R^3 and R^4 are as defined in claim 1 except that any functional group is protected if necessary;

or

(d) for the preparation of those compounds of the formula I wherein R^{2a} is hydrogen, the reductive amination in the presence of a suitable reducing agent of the aldehyde or ketone of the formula VIII:



VIII

wherein Q^1, X, Y, R^1, R^{1a}, R^2, R^5, R^6 and n are as defined in claim 1 except that any functional group is protected if necessary, with an amine of the formula VII:

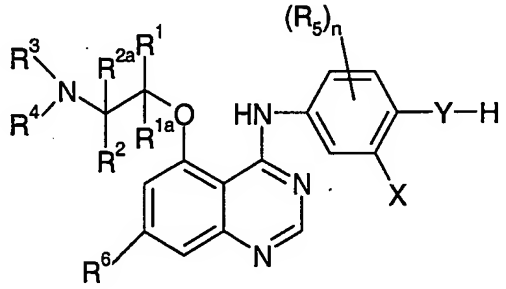
NHR^3R^4

VII

wherein R^3 and R^4 are as defined in claim 1 except that any functional group is protected if necessary;

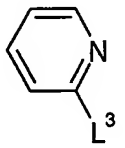
or

(e) for the preparation of those compounds of the formula **I** wherein Y is O or $N(R^7)$ and Q^1 is 2-pyridyl or 4-pyridyl the reaction, in the presence of a suitable catalyst, of a quinazoline of the formula **IV**:

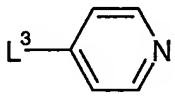


IV

wherein Y is O or $N(R^7)$ and X, R^1, R^{1a}, R^2, R^{2a}, R^3, R^4, R^5, R^6 and n are as defined in claim 1 except that any functional group is protected if necessary, with an amine of the formula **IVa** or of the formula **IVb**:



IVa

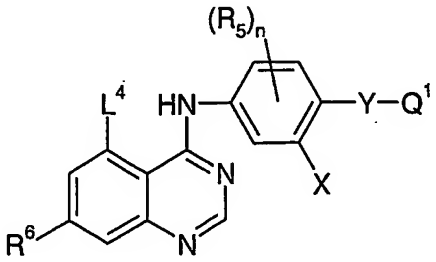


IVb

wherein L^3 is a suitable displaceable group;

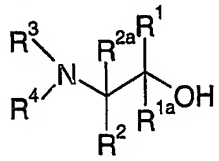
or

(f) the reaction, conveniently in the presence of a suitable phosphine and a suitable diazo compound, of a quinazoline of the formula **II**:



II

wherein R^5, R^6, Q^1, X, Y and n are as defined in claim 1 except that any functional group is protected if necessary, and L^4 is hydroxy, with an alcohol of the formula **III**:



III

wherein R^1, R^{1a}, R^2, R^{2a}, R^3 and R^4 are as defined in claim 1 except that any functional group is protected if necessary;

and thereafter, if necessary:

(i) converting a quinazoline derivative of the formula I into another quinazoline derivative of the formula I;

(ii) removing any protecting group that is present by conventional means;

(iii) forming a pharmaceutically acceptable salt.